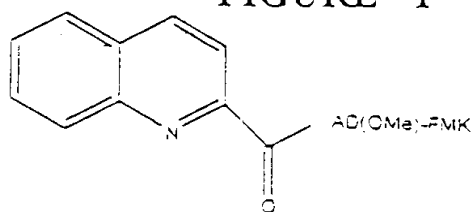
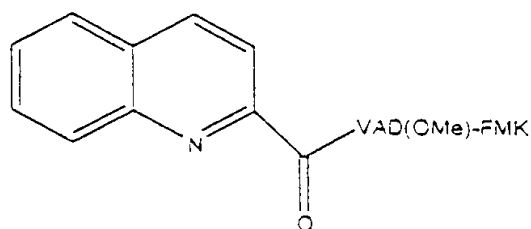


FIGURE 1



Quin-AD(OMe)-FMK M.Wt:339

FIGURE 1A



Quin-VAD(OMe)-FMK
M.Wt:483; C₂₄H₁₉N₄O₆F

FIGURE 2

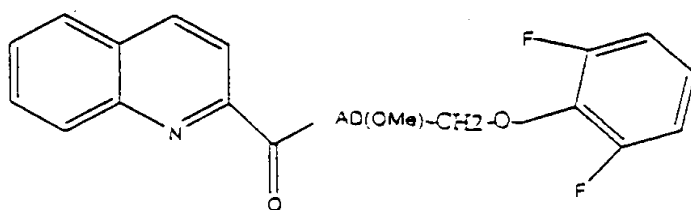
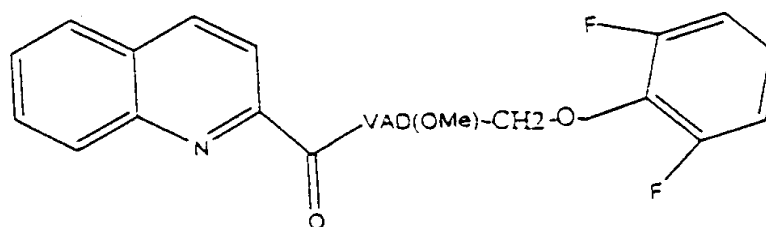


FIGURE 2A



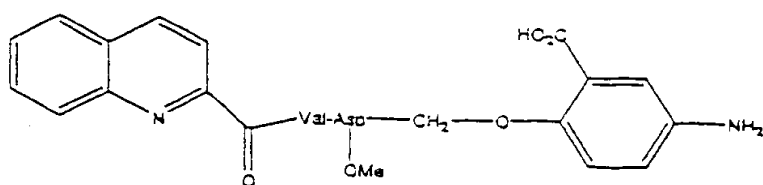


FIGURE 3

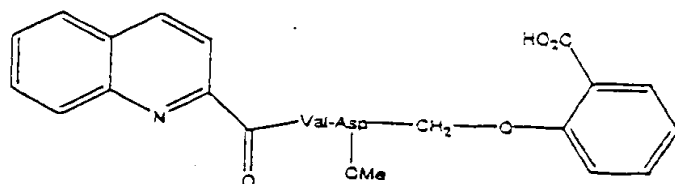


FIGURE 4

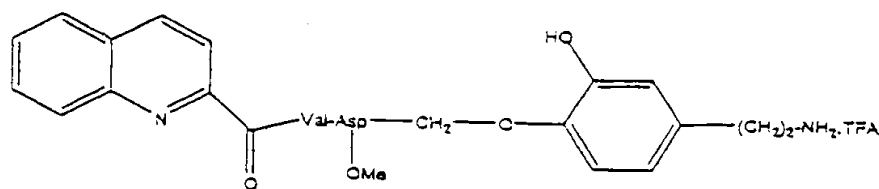


FIGURE 5

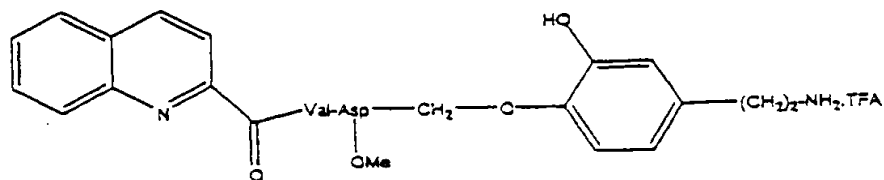


FIGURE 6

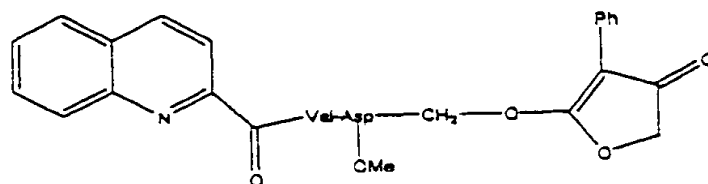


FIGURE 7

FIGURE 8

Caspase 9

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| 0.025uM | -1.602 | 0 |
| 0.05uM | -1.301 | 0 |
| 0.1uM | -1 | 0 |
| 0.5uM | -0.301 | 0 |
| 1uM | 0 | 16.2 |
| 2.5uM | 0.3979 | 21.8 |
| 5uM | 0.6989 | 47.4 |
| 10uM | 1 | 62 |
| 25uM | 1.398 | 82.4 |
| 50uM | 1.6989 | 92.6 |

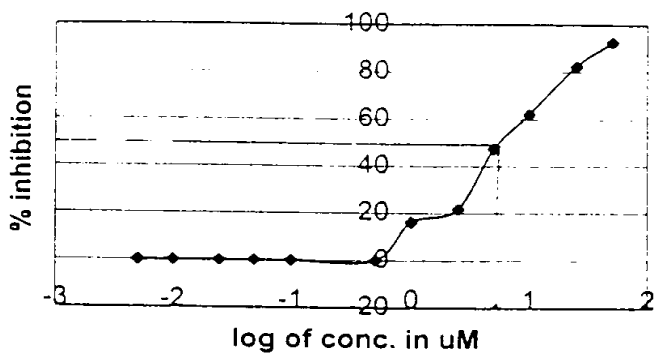
Q-(C=O)-VD(OMe)-CH₂-ASA

FIGURE 9

Caspase 8

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| 0.025uM | -1.602 | 0 |
| 0.05uM | -1.301 | 0 |
| 0.1uM | -1 | 0 |
| 0.5uM | -0.301 | 4.7 |
| 1uM | 0 | 5.5 |
| 2.5uM | 0.3979 | 21.1 |
| 5uM | 0.6989 | 45.5 |
| 10uM | 1 | 73.6 |
| 25uM | 1.398 | 96.8 |
| 50uM | 1.6989 | 99.8 |

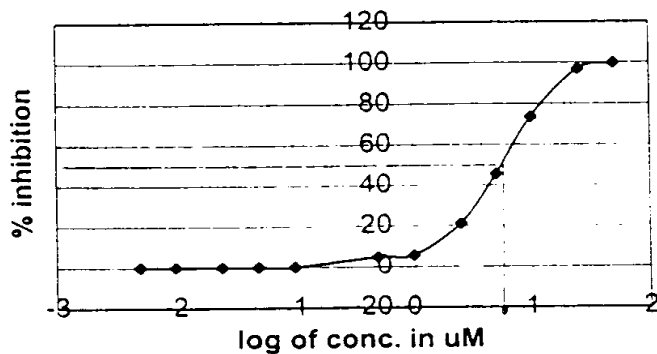
Q-(C=O)-VD(OMe)-CH₂-ASA

FIGURE 10

Caspase 1

| inh conc | log of con | % inhib |
|----------|------------|---------|
| .025uM | -1.602 | 0 |
| .05uM | -1.301 | 0 |
| .1uM | -1 | 0 |
| 0.5uM | -0.301 | 18.2 |
| 1uM | 0 | 34.8 |
| 2.5uM | 0.3979 | 69.7 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |

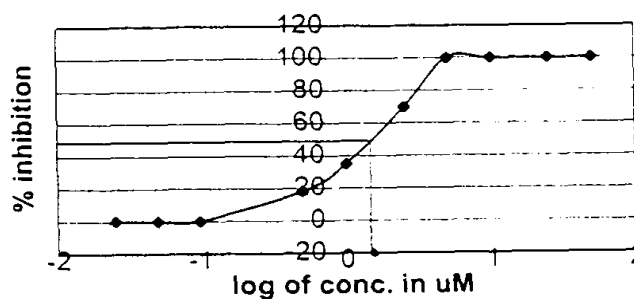
Q-(C=O)-VD(OMe)-CH₂-ASA

FIGURE 11

Caspase 3

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| .025uM | -1.602 | 2.3 |
| .05uM | -1.301 | 9.1 |
| .1uM | -1 | 6.4 |
| 0.5uM | -0.301 | 29.3 |
| 1uM | 0 | 45 |
| 2.5uM | 0.3979 | 74.8 |
| 5uM | 0.6989 | 91.5 |
| 10uM | 1 | 98.2 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |

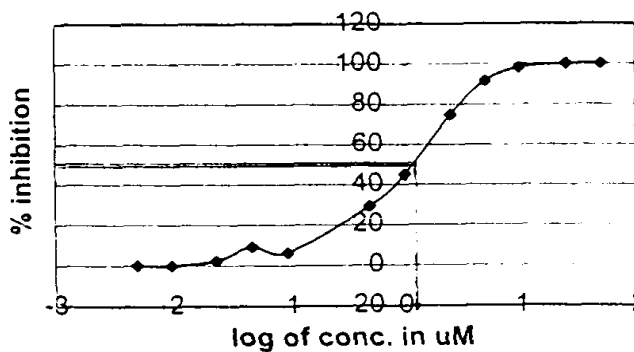
Q-(C=O)-VD(OMe)-CH₂-ASA

FIGURE 12

Caspase 1

| inh conc | log of con | % inhib |
|----------|------------|---------|
| .0025uM | -2.602 | 3.14 |
| .005uM | -2.301 | 2.6 |
| .01uM | -2 | 1.4 |
| .025uM | -1.602 | 10.3 |
| .05uM | -1.301 | 8.3 |
| .1uM | -1 | 23.7 |
| 0.5uM | -0.301 | 50.9 |
| 1uM | 0 | 66.29 |
| 2.5uM | 0.3979 | 90.3 |
| 5uM | 0.6989 | 96.3 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

Indole-(C=O)-VD(OMe)-CH₂-OPh

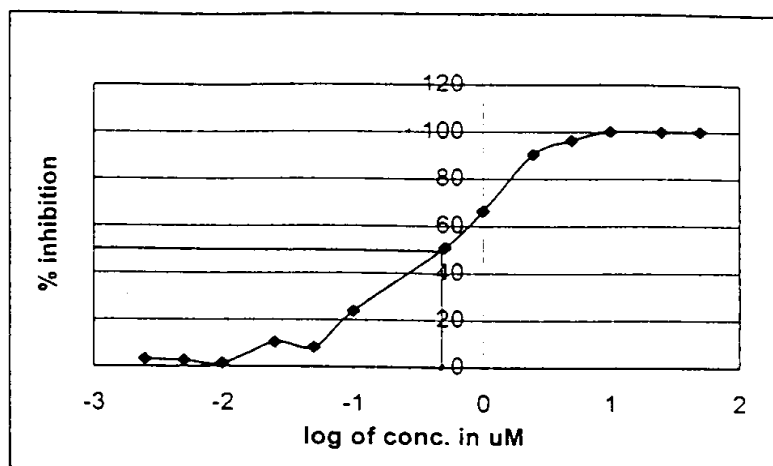


FIGURE 13

Caspase 1

| inh conc | log of con | % inhib |
|----------|------------|---------|
| .0025uM | -2.602 | 16.3 |
| .005uM | -2.301 | 19.4 |
| .01uM | -2 | 22.6 |
| .025uM | -1.602 | 42.86 |
| .1uM | -1 | 44 |
| 0.5uM | -0.301 | 74 |
| 1uM | 0 | 87.4 |
| 2.5uM | 0.3979 | 97.1 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

Melatonin-VD(OMe)-CH₂-OPh

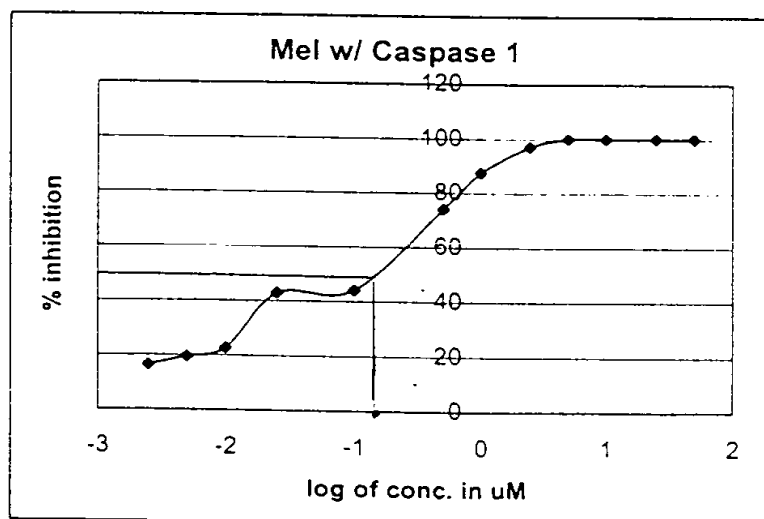


FIGURE 14

| Caspase 1 | | |
|-----------|------------|---------|
| inh conc | log of con | % inhib |
| 0.0025uM | -2.602 | 0 |
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| .025uM | -1.602 | 0 |
| .05uM | -1.301 | 7.3 |
| .1uM | -1 | 26.8 |
| 0.5uM | -0.301 | 93.4 |
| 1uM | 0 | 99.6 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |

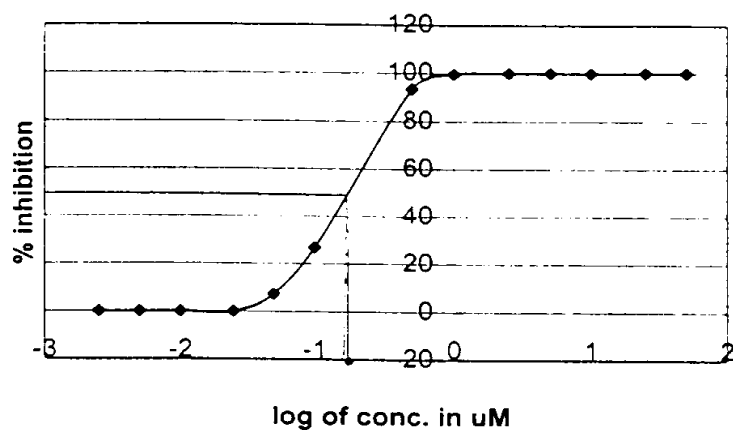
Bzl-Melatonin-VD(OMe)-CH₂-OPh

FIGURE 15

| Caspase 1 | | |
|-----------|------------|---------|
| inh conc | log of con | % inhib |
| 0.0025uM | -2.602 | 38.4 |
| 0.005uM | -2.301 | 25.7 |
| 0.01uM | -2 | 29.6 |
| .025uM | -1.602 | 23 |
| .05uM | -1.301 | 44.3 |
| 0.5uM | -0.301 | 57.2 |
| 1uM | 0 | 91.4 |
| 2.5uM | 0.3979 | 95 |
| 5uM | 0.6989 | 96.9 |

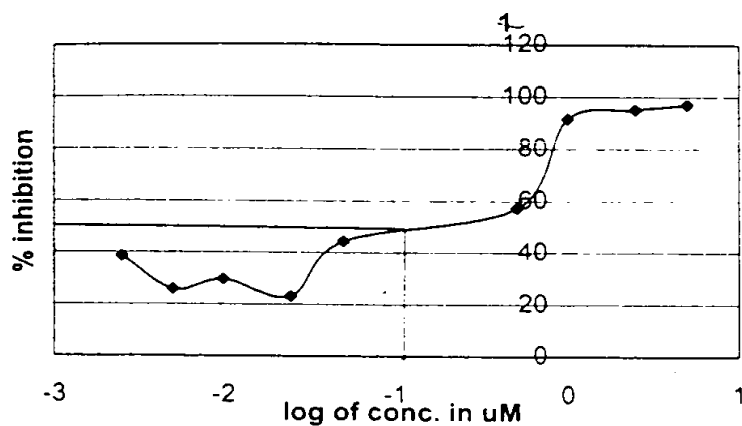
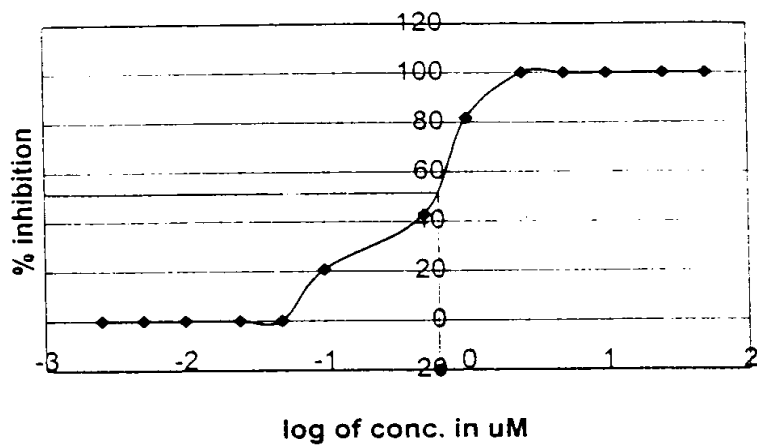
HydroxyTryptophan-VD(OMe)-CH₂-OPh

FIGURE 16

Caspase 1

TRP-VD(OCH₃)-CH₂-OPh · TFA

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.0025uM | -2.602 | 0 |
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| 0.025uM | -1.602 | 0 |
| 0.05uM | -1.301 | 0 |
| 0.1uM | -1 | 20.7 |
| 0.5uM | -0.301 | 42.7 |
| 1uM | 0 | 81.7 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |



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FIGURE 17A

Caspase 9

Q-(C=O)-L-D-(OMe)-CH₂-F (the FMK)

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 33.6 |
| 0.5uM | -1.301 | 43.9 |
| 1uM | -1 | 58.7 |
| 0.5uM | -0.301 | 90.7 |
| 1uM | 0 | 94.7 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

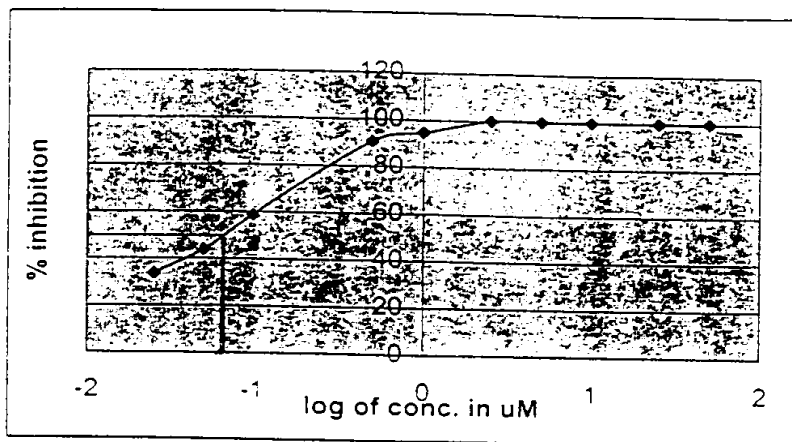


FIGURE 17B

Caspase 9

Q-(C=O)-L-D-(OMe)-CH₂-F (the FMK)

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 25.7 |
| 0.5uM | -1.301 | 37.3 |
| 1uM | -1 | 58.9 |
| 0.5uM | -0.301 | 88.9 |
| 1uM | 0 | 94.9 |
| 2.5uM | 0.3979 | 96.1 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

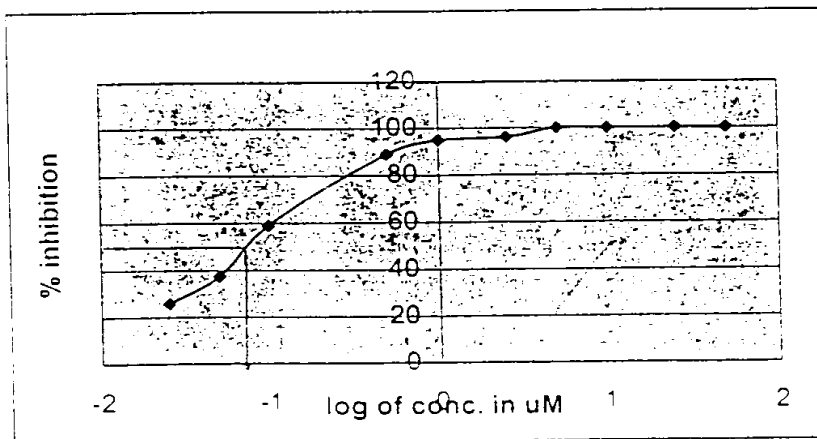


FIGURE 18A

Caspase 9

 $Q-(C=O)-V-D-(OCH_3)-CH_2-F$ (the FMK)

| conc | log of conc | % inhib |
|---------|-------------|---------|
| 0.025uM | -1.602 | 47.3 |
| 0.05uM | -1.301 | 64.4 |
| 0.1uM | -1 | 81.2 |
| 0.5uM | -0.301 | 97.8 |
| 1uM | 0 | 99.5 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

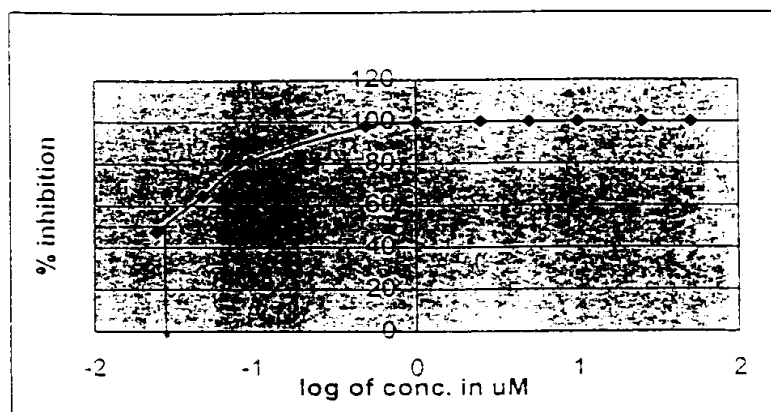


FIGURE 18B

Caspase 9

 $Q-(C=O)-V-D-(OCH_3)-CH_2-F$ (the FMK)

| conc | log of conc | % inhib |
|---------|-------------|---------|
| 0.025uM | -1.602 | 62.2 |
| 0.05uM | -1.301 | 75.3 |
| 0.1uM | -1 | 81.3 |
| 0.5uM | -0.301 | 99.1 |
| 1uM | 0 | 100 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

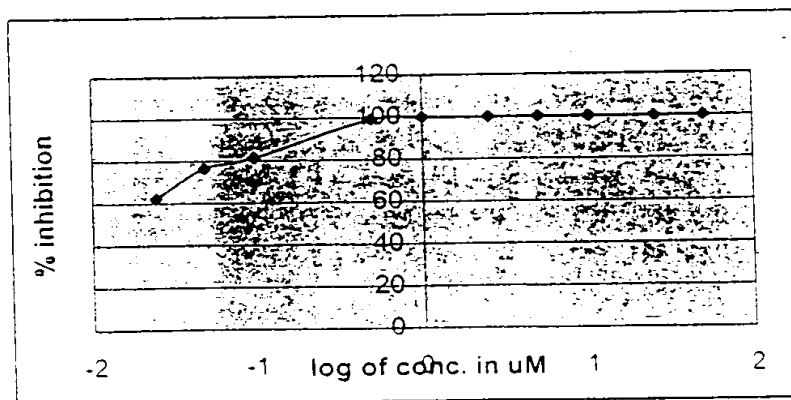


FIGURE 19

Caspase 1

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.0025uM | -2.602 | 0 |
| 0.005uM | -2.301 | 3.2 |
| 0.01uM | -2 | 0 |
| .025uM | -1.602 | 1.1 |
| .05uM | -1.301 | 0 |
| .1uM | -1 | 0 |
| 0.5uM | -0.301 | 15.3 |
| 1uM | 0 | 33.7 |
| 2.5uM | 0.3979 | 72.1 |
| 5uM | 0.6989 | 97.4 |
| 10uM | 1 | 100 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |

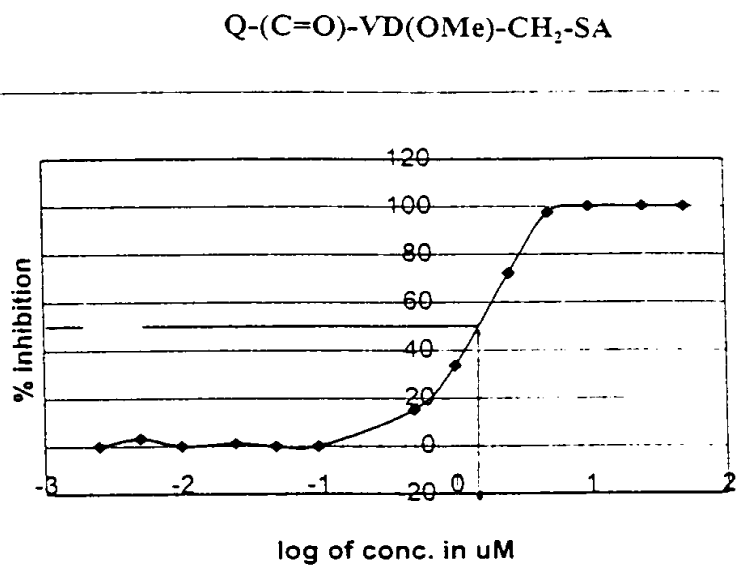


FIGURE 20

Caspase 3

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.005uM | -2.301 | 0 |
| 0.01uM | -2 | 0 |
| .025uM | -1.602 | 0.57 |
| .05uM | -1.301 | 2.8 |
| .1uM | -1 | 18.3 |
| 0.5uM | -0.301 | 32.4 |
| 1uM | 0 | 54.7 |
| 2.5uM | 0.3979 | 87.8 |
| 5uM | 0.6989 | 97.6 |
| 10uM | 1 | 99.7 |
| 25uM | 1.398 | 100 |
| 50uM | 1.6989 | 100 |

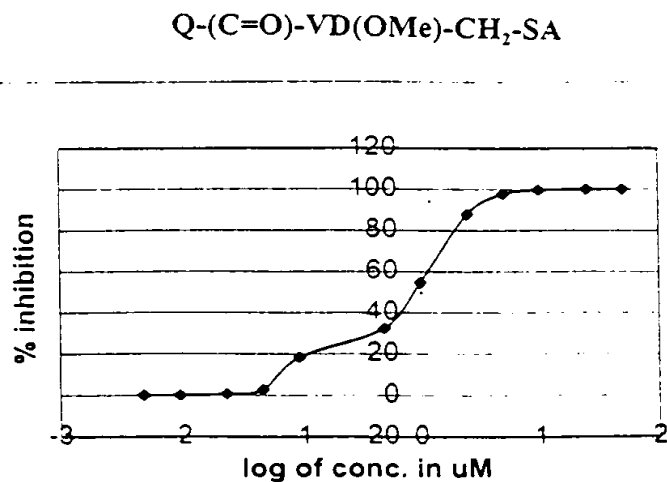


FIGURE 21



Caspase 1

| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 19 |
| 0.5uM | -1.301 | 22 |
| 1uM | -1 | 19 |
| 0.5uM | -0.301 | 46.7 |
| 1uM | 0 | 69.5 |
| 2.5uM | 0.3979 | 92.7 |
| 5uM | 0.6989 | 98.5 |
| 10uM | 1 | 87.3 |

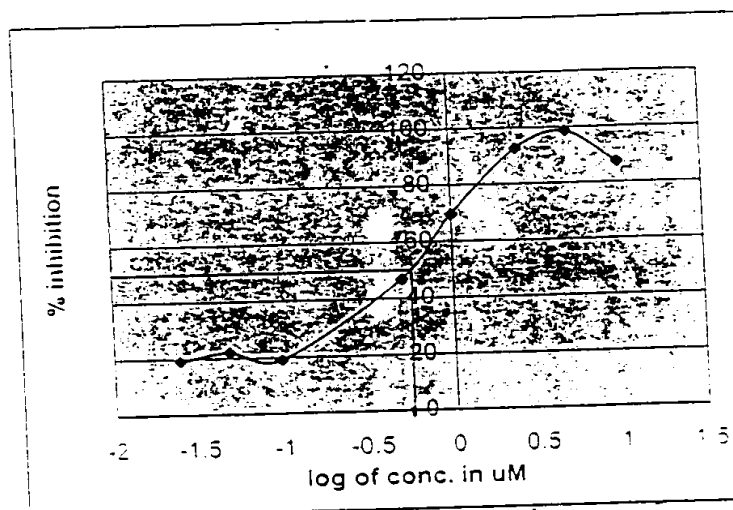
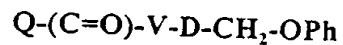
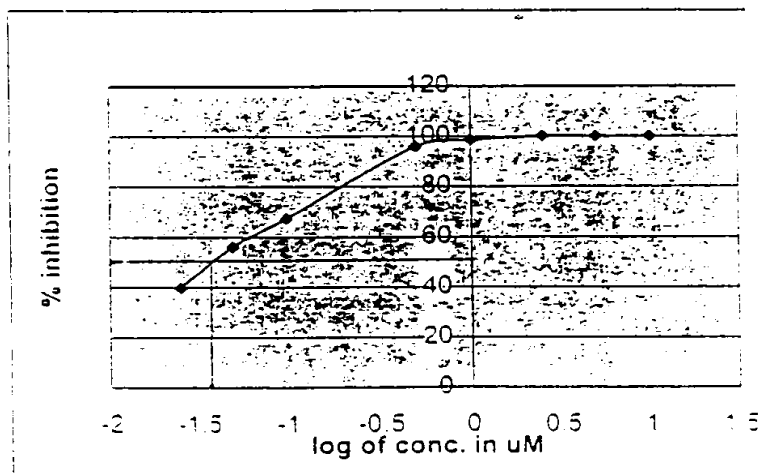


FIGURE 22



Caspase 1

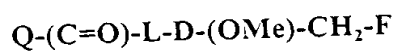
| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 39.8 |
| 0.5uM | -1.301 | 55.98 |
| 1uM | -1 | 67.2 |
| 0.5uM | -0.301 | 95.8 |
| 1uM | 0 | 98.5 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |



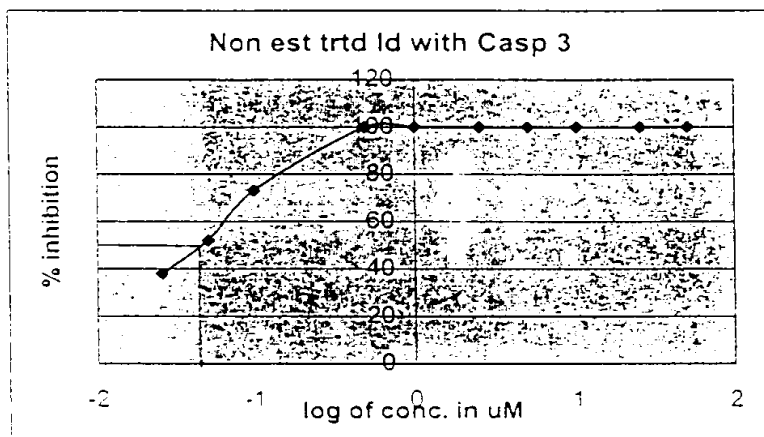
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FIGURE 25A

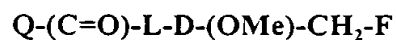
Non-esterase treated, Inhibitor D with Caspase 3



| inh conc | log of conc | % inhib |
|----------|-------------|---------|
| 0.025uM | -1.602 | 37.8 |
| 0.05uM | -1.301 | 52 |
| 0.1uM | -1 | 73 |
| 0.5uM | -0.301 | 100 |
| 1uM | 0 | 100 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

**FIGURE 25B**

Esterase treated, Inhibitor D with Caspase 3



| inh conc | log of conc | % inhib |
|----------|-------------|---------|
| 0.025uM | -1.602 | 38.2 |
| 0.05uM | -1.301 | 68.9 |
| 0.1uM | -1 | 80.7 |
| 0.5uM | -0.301 | 97.6 |
| 1uM | 0 | 96.6 |
| 2.5uM | 0.3979 | 96.2 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

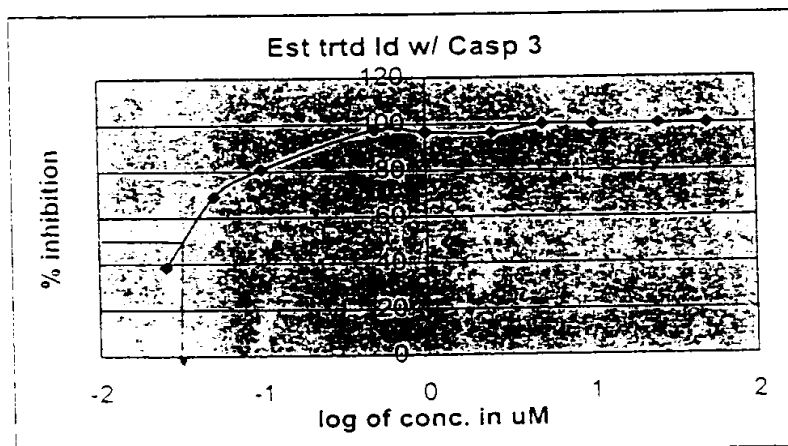
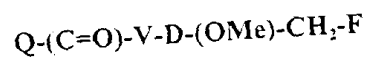


FIGURE 23

Esterase treated Inhibitor C with Caspase 1



| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 40.1 |
| 0.5uM | -1.301 | 54.9 |
| 1uM | -1 | 73.2 |
| 0.5uM | -0.301 | 81.7 |
| 1uM | 0 | 100 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

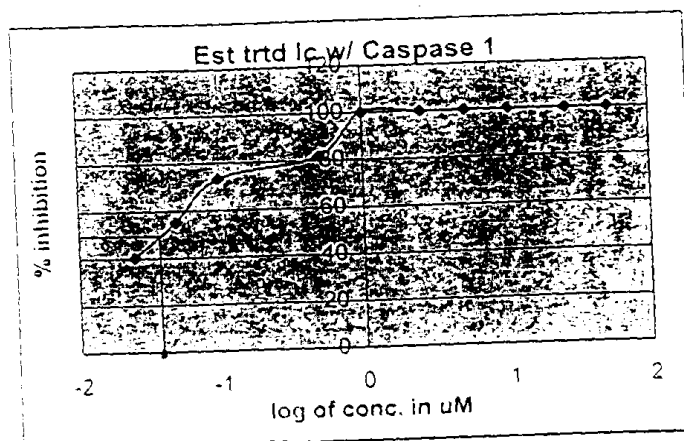
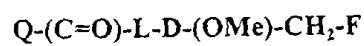


FIGURE 24

Esterase treated Inhibitor D with Caspase 1



| inh conc | log of con | % inhib |
|----------|------------|---------|
| 0.25uM | -1.602 | 0 |
| 0.5uM | -1.301 | 33.8 |
| 1uM | -1 | 63.4 |
| 0.5uM | -0.301 | 85.2 |
| 2.5uM | 0.3979 | 85.2 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.6979 | 100 |

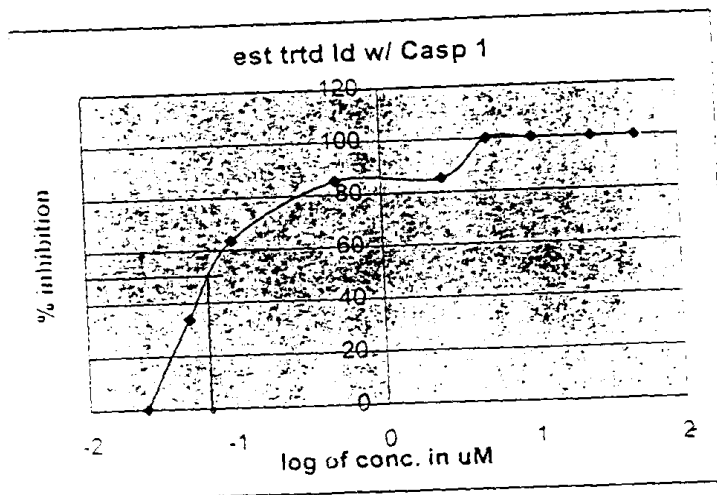


FIGURE 26

Q-LD-OPh

| | | |
|---------|--------|------|
| 0.05 uM | -1.301 | 5.5 |
| 0.1 uM | -1 | 11 |
| 0.5 uM | -0.301 | 46 |
| 1 uM | 0 | 68 |
| 2.5 uM | 0.3979 | 86.8 |
| 5 uM | 0.6989 | 94.5 |
| 10 uM | 1 | 100 |
| 25 uM | 1.3979 | 100 |
| 50 uM | 1.6989 | 100 |

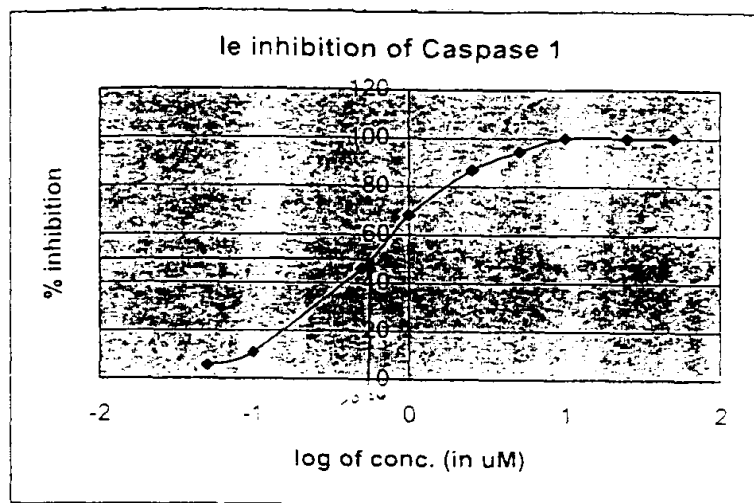
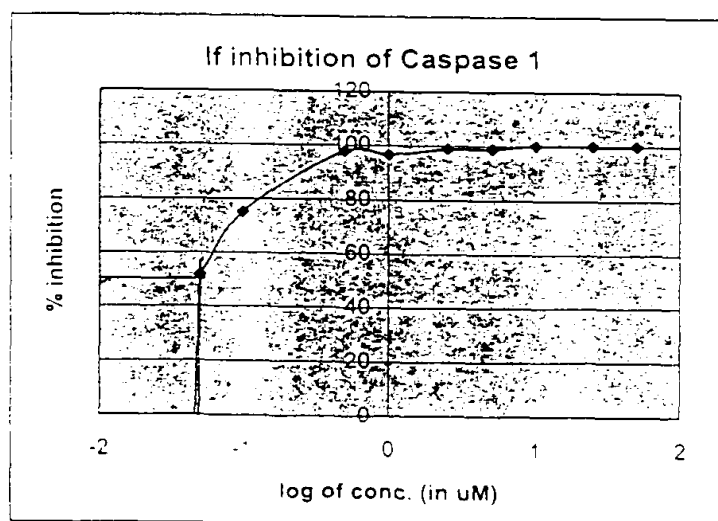


FIGURE 27

Q-VD-OPh

IC₅₀ [uM] % inhibition

| | | |
|---------|--------|------|
| 0.05 uM | -1.301 | 51.6 |
| 0.1 uM | -1 | 75 |
| 0.5 uM | -0.301 | 97.8 |
| 1 uM | 0 | 96.7 |
| 2.5 uM | 0.3979 | 98.9 |
| 5 uM | 0.6989 | 98.9 |
| 10 uM | 1 | 100 |
| 25 uM | 1.3979 | 100 |
| 50 uM | 1.6989 | 100 |



Caspase 3 w/ IE

Q-(C=O)-LD-CH₂-O-Ph

| inh conc | log of con | % inhib |
|----------|------------|---------|
| .025uM | -1.602 | 31.85 |
| .05uM | -1.301 | 47.1 |
| .1uM | -1 | 59.2 |
| 0.5uM | -0.301 | 96.2 |
| 1uM | 0 | 100 |
| 2.5uM | 0.3979 | 100 |
| 5uM | 0.6989 | 100 |
| 10uM | 1 | 100 |
| 25uM | 1.3979 | 100 |
| 50uM | 1.699 | 100 |

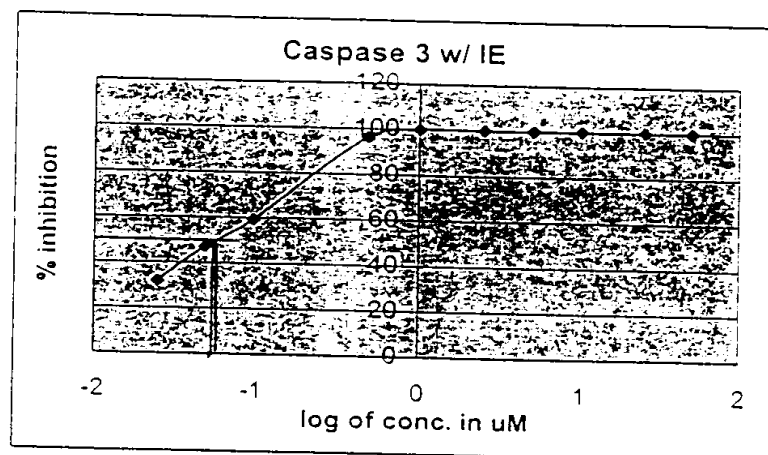


FIGURE 28

FIGURE 29

IMPORTANT AMINO ACIDS

